CLAIMS

What is claimed is:

1. A method of treating disease resulting from malformed proteins from a mammal comprising:

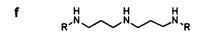
administering to said mammal a therapeutically effective amount of a bis-cyclic compound;

wherein said bis-cyclic compound is characterized by clearing malformed proteins and by an ability to cross a blood brain barrier of said mammal.

2. The method of claim 1, wherein the compound is comprised of a linking group which covalently binds together two cyclic moieties having the general structural formula I:

wherein each of positions 1-9 may be independently substituted.

3. The method of claim 2, wherein the linking group is chosen from



$$g \qquad {}_{R}, \overset{H}{N} \searrow \overset{H}{N} \searrow \overset{R}{N} \stackrel{R}{\searrow} \overset{R}{\searrow}$$

$$i$$
 R^{N}
 N
 N
 N
 N
 N

$$\mathbf{m} \qquad \mathbf{R}^{\mathsf{N}} \searrow \mathbf{0} \searrow \mathbf{0} \searrow \mathbf{N}^{\mathsf{N}}$$

$$\mathbf{n}$$
 \mathbf{R} \mathbf{N} \mathbf{N} \mathbf{N} \mathbf{N}

$$\mathbf{p}$$
 $R_1 = \mathbf{N} \mathbf{H}_2$

$$\mathbf{u} = \mathbf{R}_2 - \mathbf{N}_2$$

wherein each "R" is independently any moiety of formula I.

4. The method of claim 3, wherein each "R" is independently chosen from

- 5. The method of claim 3, wherein each "R" is quinacrine.
- 6. The method of Claim 1, wherein said mammal is selected from the group consisting of a human, cow, pig, sheep and goat.

7. The method of Claim 1, wherein a position chosen from positions 1-9 of formula I is substituted with a moiety chosen from

8. The method of Claim 1, wherein a position chosen from position 1-9 of formula I is substituted with a moiety chosen from

9. The method of Claim 1, wherein the malformed protein and its associated disease is selected from the group consisting of:

Disease	Insoluble	Proteins
<u>Disease</u>	IIISOIUUIE	Proteins

Alzheimer's Disease APP, Aß peptide, al-antichymotrypsin,

tan, non-Aß component

Prion diseases, Creutzfeld Jakob disease, scrapie and bovine spongeform

Encephalopathy PrP^{Sc}

ALS SOD and neurofilament

Pick's disease Pick body

Parkinson's disease Lewy body

Diabetes Type 1 Amylin

Multiple myeloma-- IgGL-chain plasma cell dyscrasias

Familial amyloidotic Transthyretin

polyneuropathy

Medullary carcinoma of

thyroid

Procalcitonin

Chronic renal failure

 β_2 --microglobulin

Congestive heart failure

Atrial natriuretic factor

Senile cardiac and

systemic amyloidosis

Transthyretin

Chronic inflammation

Serum amyloid A

Atherosclerosis

ApoA1

Familial amyloidosis

Gelsolin.

The method of Claim 1, wherein the disease and its associated malformed prion is 10. selected from the group consisting of

Alzheimer's Disease

APP, Aβ peptide, α1-

antichymotrypsin, tan, non-

Aβ component

Prion diseases, Creutzfeld Jakob disease, scrapie and

bovine spongeform

PrPSc Encephalopathy

Parkinson's disease

Lewy body

Diabetes Type 1

Amylin

Familial amyloidotic

polyneuropathy

Transthyretin.

- 11. The method according to Claim 8, wherein the oral administration step is in an amount of about 100 mg to 10,000 mg/day/75 kg of body weight.
- 12. The method of Claim 1, wherein the administration step comprises administration by injection.

- The method of Claim 1, wherein the administration step comprises a technique selected from the group consisting of transdermal administration, subcutaneous injection, intravenous injection, intraperitoneal injection, intramuscular injection, intrasternal injection, intrathecal injection, intranasal, and infusion techniques.
- 14. The method as claimed in Claim 5, wherein the quinacrine is 100% dextrorotary quinacrine.
- 15. The method of Claim 5, wherein the mammal is suffering from Creutzfeldt-Jakob disease.
- The method of Claim 5, wherein the mammal is suffering from a disease selected from the group consisting of scrapie, transmissible spongioform encephalopathy (TSE), Alzheimer's disease, Parkinson's disease, amyotrophic lateral sclerosis, multiple sclerosis, autism, schizophrenia, bipolar disorders, fronto-temporal dementia, Pick's disease, progressive supranuclear palsy, diffuse Lewy body disease, systemic lupus erythematosus, rheumatoid arthritis, Huntington's disease, spinocerebellar ataxias, diabetes mellitus, Types I and II, Crohn's disease, ulcerative colitis, systemic amyloidosis, primary amyloidosis, polyneuropathy and AIDS.
- 17. A composition for treating livestock with malformed proteins comprising: livestock feed; and a bis-cyclic compound.
- 18. The composition for treating livestock afflicted with malformed proteins as claimed in claim 17, wherein the compound is comprised of a linking group which covalently binds together two cyclic moieties having the general structural formula I:

wherein each of positions 1-9 may be independently substituted.

- 19. A method for clearing malformed proteins from livestock, said method comprising:
- a. administering a pharmaceutically effective amount of the composition of Claim 17; and
- b. repeatedly providing said livestock feed to livestock over a therapeutically effective period of time.
- 20. A method for clearing malformed proteins from livestock, said method comprising:
- a. administering a pharmaceutically effective amount of the composition of Claim 18; and
- b. repeatedly providing said livestock feed to livestock over a therapeutically effective period of time.
- 21. A composition, comprising:

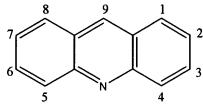
livestock feed; and

a bis-compound comprising two tricyclic moieties covalently bound by a linking group.

The composition of claim 21, wherein both of the tricyclic moieties are quinacrine.

23. The composition of Claim 21, wherein the tricyclic moiety is chosen from

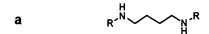
- 24. A composition, comprising:
 - a pharmaceutically acceptable carrier; and
- a bis-cyclic compound, comprised of two cyclic moieties covalently bound together by a linking group, wherein the cyclic moieties have the general structural formula I



Acridine

wherein each of positions 1-9 may be independently substituted.

25. The composition of claim 24, wherein the linking group is chosen from



$$d = R^{-N} \longrightarrow N^{-R}$$

- t R_2 $\stackrel{H}{\sim}$ $\stackrel{H}{\sim}$ $\stackrel{N}{\sim}$ $\stackrel{N}{\sim}$ $\stackrel{R_2}{\sim}$
- u R₂ N N R₂
- 26. A compound chosen from bis-(6-chloro-2-methoxy-acridin-9-yl) and an analog thereof.
- 27. A compound chosen from bis-(7-chloro-2-methoxy-benzo[b][1,5]naphthyridin-10-yl) and an analog thereof
- A compound chosen from (6-chloro-2-methoxy-acridin-9-yl)-(3-{4-[3-(6-chloro-2-methoxy-acridin-9-ylamino)-propyl]-piperazin-1-yl}-propyl)-amine, *N,N*'-bis-(6-chloro-2-methoxy-acridin-9-yl)-1,8-diamino-3,6-dioxaoctane, and (1-{[4-(6-chloro-2-methoxy-acridin-9-ylamino)-propyl]-carbamoyl}-ethyl)-carbamic acid *tert*-butyl ester.